

GEPHE SUMMARY

SCN8A (Nav1.6) (https://www.gephebase.org/search-criteria?/and+Gene) Gephebase= [^] SCN8A (Nav1.6) [^] #gephebase-summary-title	Gephebase Gene	GP00001655	GepheID
Published	Entry Status	Prigent	Main curator

PHENOTYPIC CHANGE

Physiology (https://www.gephebase.org/search-criteria?/and+Trait) Category= [^] Physiology [^] #gephebase-summary-title	Trait Category		
Xenobiotic resistance (TTX) (https://www.gephebase.org/search-criteria?/and+Trait) = [^] Xenobiotic resistance (TTX) [^] #gephebase-summary-title	Trait		
TTX-Sensitive Erythrolamprus (=Liophis) poecilogyrus	Trait State in Taxon A		
TTX-Resistant Erythrolamprus (=Liophis) epinephelus	Trait State in Taxon B		
	Ancestral State		
Taxon A	Taxonomic Status		
Interspecific (https://www.gephebase.org/search-criteria?/and+Taxonomic) Status= [^] Interspecific [^] #gephebase-summary-title			
	Taxon A		Taxon B
Erythrolamprus poecilogyrus (https://www.gephebase.org/search-criteria?/and+Taxon and Synonyms= [^] Erythrolamprus poecilogyrus [^] #gephebase-summary-title)	Latin Name	Erythrolamprus epinephelus (https://www.gephebase.org/search-criteria?/and+Taxon and Synonyms= [^] Erythrolamprus epinephelus [^] #gephebase-summary-title)	Latin Name
-	Common Name	-	Common Name
	Synonyms		Synonyms
Coluber poecilogyrus; Liophis poecilogyrus; Coluber poecilogyrus Wied-Neuwied, 1825; Liophis poecilogyrus (Wied-Neuwied, 1825); AMNH 3594; AMNH:3594		Erythrolamprus epinephelus; Leimadophis epinephelus; Liophis epinephelus; Liophis epinephelus Cope, 1862; ANSP 3688; ANSP:3688	
species	Rank	species	Rank
	Lineage		Lineage
cellular organisms; Eukaryota; Opisthokonta; Metazoa; Eumetazoa; Bilateria; Deuterostomia; Chordata; Craniata; Vertebrata; Gnathostomata; Teleostomi; Euteleostomi; Sarcopterygii; Dipnotetrapodomorpha; Tetrapoda; Amniota; Sauropsida; Sauria; Lepidosauria; Squamata; Bifurcata; Unidentata; Episquamata; Toxicofera; Serpentes; Colubroidea; Dipsadidae; Erythrolamprus		cellular organisms; Eukaryota; Opisthokonta; Metazoa; Eumetazoa; Bilateria; Deuterostomia; Chordata; Craniata; Vertebrata; Gnathostomata; Teleostomi; Euteleostomi; Sarcopterygii; Dipnotetrapodomorpha; Tetrapoda; Amniota; Sauropsida; Sauria; Lepidosauria; Squamata; Bifurcata; Unidentata; Episquamata; Toxicofera; Serpentes; Colubroidea; Dipsadidae; Erythrolamprus	
Erythrolamprus () - (Rank: genus) (https://www.ncbi.nlm.nih.gov/Taxonomy/Browser/wwwtax.cgi?id=121327)	Parent	Erythrolamprus () - (Rank: genus) (https://www.ncbi.nlm.nih.gov/Taxonomy/Browser/wwwtax.cgi?id=121327)	Parent
338838 (https://www.ncbi.nlm.nih.gov/Taxonomy/Browser/wwwtax.cgi?id=338838)	NCBI Taxonomy ID	758879 (https://www.ncbi.nlm.nih.gov/Taxonomy/Browser/wwwtax.cgi?id=758879)	NCBI Taxonomy ID
No	is Taxon A an Intraspecies?	No	is Taxon B an Intraspecies?

GENOTYPIC CHANGE

SCN8A	Generic Gene Name	UniProtKB Erythrolamprus epinephelus A0A1B0Z7B3 (http://www.uniprot.org/uniprot/A0A1B0Z7B3)
-	Synonyms	0
-	String	
	Sequence Similarities	
Belongs to the sodium channel (TC 1.A.1.10) family.		
GO:0005244 : voltage-gated ion channel activity (https://www.ebi.ac.uk/QuickGO/term/GO:0005244)	GO - Molecular Function	
GO:0005248 : voltage-gated sodium channel activity (https://www.ebi.ac.uk/QuickGO/term/GO:0005248)		
GO:0034765 : regulation of ion transmembrane transport (https://www.ebi.ac.uk/QuickGO/term/GO:0034765)	GO - Biological Process	

GO:0001518 : voltage-gated sodium channel complex
 (<https://www.ebi.ac.uk/QuickGO/term/GO:0001518>)

No (<https://www.gephebase.org/search-criteria?/and+Presumptive Null=^No^#gephebase-summary-title>)

Coding (<https://www.gephebase.org/search-criteria?/and+Molecular Type=^Coding^#gephebase-summary-title>)

SNP (<https://www.gephebase.org/search-criteria?/and+Aberration Type=^SNP^#gephebase-summary-title>)

Nonsynonymous

G1717M in DIV domain (not tested)

Candidate Gene (<https://www.gephebase.org/search-criteria?/and+Experimental Evidence=^Candidate Gene^#gephebase-summary-title>)

Presumptive Null

Molecular Type

Aberration Type

SNP Coding Change

Molecular Details of the Mutation

Experimental Evidence

	Taxon A	Taxon B	Position
Codon	-	-	-
Amino-acid	-	-	-

Main Reference

Historical Contingency in a Multigene Family Facilitates Adaptive Evolution of Toxin Resistance. (2016) (<https://pubmed.ncbi.nlm.nih.gov/27291053>)

Authors

McGlothlin JW; Kobiela ME; Feldman CR; Castoe TA; Geffney SL; Hanifin CT; Toledo G; Vonk FJ; Richardson MK; Brodie ED; Pfrender ME; Brodie ED

Abstract

Novel adaptations must originate and function within an already established genome [1]. As a result, the ability of a species to adapt to new environmental challenges is predicted to be highly contingent on the evolutionary history of its lineage [2-6]. Despite a growing appreciation of the importance of historical contingency in the adaptive evolution of single proteins [7-11], we know surprisingly little about its role in shaping complex adaptations that require evolutionary change in multiple genes. One such adaptation, extreme resistance to tetrodotoxin (TTX), has arisen in several species of snakes through coevolutionary arms races with toxic amphibian prey, which select for TTX-resistant voltage-gated sodium channels (Nav) [12-16]. Here, we show that the relatively recent origins of extreme toxin resistance, which involve the skeletal muscle channel Nav1.4, were facilitated by ancient evolutionary changes in two other members of the same gene family. A substitution conferring TTX resistance to Nav1.7, a channel found in small peripheral neurons, arose in lizards \sim 4170 million years ago (mya) and was present in the common ancestor of all snakes. A second channel found in larger myelinated neurons, Nav1.6, subsequently evolved resistance in four different snake lineages beginning \sim 438 mya. Extreme TTX resistance has evolved at least five times within the past 12 million years via changes in Nav1.4, but only within lineages that previously evolved resistant Nav1.6 and Nav1.7. Our results show that adaptive protein evolution may be contingent upon enabling substitutions elsewhere in the genome, in this case, in paralogs of the same gene family.

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Additional References

RELATED GEPHE

Related Genes

1 (SCN4A (Nav1.4)) (<https://www.gephebase.org/search-criteria?/or+Taxon ID=^338838^/and+Trait=Xenobiotic resistance/or+Taxon ID=^758879^/and+Trait=Xenobiotic resistance/and+groupHaplotypes=true#gephebase-summary-title>)

Related Haplotypes

No matches found.

EXTERNAL LINKS

COMMENTS

Non-null mutation